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African-American and caucasian women with newly diagnosed breast cancer were enrolled in the study. Upon enrollment, demographic data, tumor characteristics, and tumor staging data were obtained. Tumor samples were evaluated for estrogen and progesterone receptor levels, DNA ploidy, S-phase fraction, HER-2/neu expression, p53 protein accumulation, cathepsin D levels, and glutathione levels. Treatment choices, treatment received, relapse site and date, and date and cause of death were recorded in follow-up. To date, 140 women have been enrolled in the study of tumor biology and 290 women have been evaluated in the clinical presentation and treatment phases of the study. No racial differences in clinical presentation, tumor stage, tumor biologic characteristics, or treatment have been identified. In addition, there has been no difference in time to treatment failure or overall survival. We conclude that economically similar black and white women with breast cancer are likely to have similar clinical manifestations and outcomes.

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Introduction

Breast cancer is the most common invasive malignancy affecting American women, accounting for 28% of all tumors diagnosed in this group. It is also a leading cause of cancer related death in the United States. Although the age-adjusted incidence of breast cancer in black women in the U.S. is less than that seen in white women, the mortality rates observed in blacks and whites are virtually identical. This discrepancy is the result of a significantly lower five year survival rate for black women when compared to white women with breast cancer. The most recent results from the Surveillance, Epidemiology, and End Results (SEER) program of the National Cancer Institute have documented an 80% five year relative survival for white women diagnosed with breast cancer between 1983 and 1989; the corresponding rate for black women was only 64%. While improvements in the detection and treatment of breast cancer over the last 30 years have led to an improved five year relative survival, there is no evidence that these advances have had an influence on the racial differences in survival.

In order to improve the survival of black women with breast cancer, an understanding of the factors which contribute to their poorer prognosis is necessary. It is known that black women generally have more advanced disease than white women at the time of initial presentation. A tendency toward larger primary tumors as well as a lower incidence of disease confined to the breast and a higher incidence of distant metastases at the time of diagnosis have been documented. 3-13 Environmental, behavioral, and biological factors have also been used to explain the higher incidence of advanced disease in black women with breast cancer. In particular, attention has focused on issues relating to access to medical care and preventive health services. The use of screening mammography has not been shown to be significantly different between healthy black and white women, although the only study of racial differences in breast cancer that has addressed this issue has noted a lower incidence of prior mammography in black women with breast cancer than in their white counterparts. 12,14 Black women with breast cancer have been found to more often rely on hospital-based or public clinics for their health care. and have been noted by some investigators to have a longer interval between symptom recognition and medical consultation. 7,15-17 The difference in median time to medical consultation between black and white women has generally been short, however, and has not adequately explained the significant difference in stage of disease at presentation.

There are also several biological differences in breast cancers of black and white women which may contribute to the differences in disease stage and survival seen in these populations. There is an increased incidence of medullary carcinoma among black women, accounting for 6-9% of all breast cancers, compared to white women, where this tumor histology is seen in 2-3% of women. Black women have also been noted to have a higher incidence of poorly differentiated tumors of the breast, whether evaluated by architectural grade or nuclear grade, and in one large study higher grade tumors were significantly correlated with disease of more advanced stage. A majority of studies which have compared hormone receptor levels in black and white women have documented a lower than expected incidence of estrogen receptor positive tumors in black

women with breast cancer. ^{5,6,12,14,19-23} One group of investigators has examined some of the more recently identified markers of breast tumor biology, including DNA ploidy, Sphase fraction, HER-2/neu protein levels and p53 protein accumulation. (52) White women had a significantly lower S-phase fraction than either the black or Hispanic populations. ²⁴ This finding is not unexpected in light of the higher frequency of poorly differentiated tumors in African American women. The Black/White Cancer Survival Study, the most comprehensive study to date of racial survival differences in breast cancer found that tumor biologic characteristics (tumor grade and hormone receptor status) were second only to tumor stage in contributing to the observed survival difference. ²⁵

There are a limited number of studies which have evaluated the treatment of breast cancer in African American women. The Black/White Cancer Survival Study Group has reported that in women of equivalent stage, black women were just as likely to have surgical therapy as part of their primary treatment plan as white women. (73) They did find that black women were less likely to have breast conserving surgery and more likely to have a modified radical mastectomy. The use of systemic adjuvant therapy, either chemotherapy or endocrine therapy has generally not been found to vary significantly according to race, although the data in this area is quite limited. Leval Even less information is available regarding the efficacy of systemic therapy in preventing relapse or improving survival in African American women with breast cancer. One study, presented only in abstract form, suggested that black women enrolled on Eastern Cooperative Oncology Group chemotherapy studies for breast cancer had a poorer survival than matched controls, although there is not enough information presented to adequately analyze the reported findings.

Confounding all of this information is the issue of socioeconomic status and its close correlation with race. Observed differences in outcome, particularly if influenced by access to medical care, could certainly be a result of socioeconomics and not race. Attempts to control for socioeconomic factors (performed indirectly using census tract data) have not resulted in uniform agreement. Some studies have demonstrated a disappearance of racial differences in survival while others continue to show a significant impact of race upon survival with breast cancer. Notably, the only prospective study which has collected socioeconomic data from individual patients demonstrated a continued effect of race on stage of disease at presentation. In addition, the noted differences in tumor biology (histology, tumor grade, and hormone receptor status) are less easily explained solely by socioeconomic issues and thus raise the possibility of other factors significantly contributing to the observed survival difference.

We have initiated a prospective study evaluating the clinical, pathologic, and biologic characteristics of newly diagnosed breast cancers in a racially mixed, socioeconomically uniform cohort of patients seen at Truman Medical Center, the public hospital for Kansas City, Missouri. The objectives of the study are: 1) to determine if there are significant differences in breast cancer characteristics at presentation, prognostic factors, or treatment which could explain the survival differences noted between black and white women with the disease and 2) to determine if any documented differences are correlated with the survival of the women in the study.

Methods

Eligibility criteria. The study is being conducted at Truman Medical Center, the public hospital for Kansas City, Missouri. Women who meet the following eligibility requirements are being prospectively enrolled: 1) histologically confirmed invasive adenocarcinoma of the breast, 2) primary surgical therapy for the breast cancer performed at Truman Medical Center, 3) women of African-American or white ethnic background, 4) no prior exposure to radiation therapy or chemotherapy, and 5) written informed consent.

Study enrollment began in November 1991 and continues through the present time. In addition to the prospectively enrolled subjects, all women meeting the above screening criteria with their breast cancer diagnosed between January 1987 and October 1991 were included in the sections of the study dealing with clinical presentation and treatment choices, to allow for a more reliable analysis of treatment patterns.

Demographic data. After study enrollment, demographic information is obtained, including age, race, menstrual history, estrogen exposure, family history of cancer, nutritional measurements, and weekly alcohol consumption.

Tumor analysis. Tumor tissue was obtained from either breast biopsy or mastectomy specimens, after gross examination by a pathologist, and tissue was placed in zinc-buffered formalin for routine histology, frozen for routine quantitative estrogen and progesterone receptor analysis, and sent fresh for drug metabolism parameters. Hematoxylin and eosin stained sections will be examined, and the tumors classified according to the criteria of the World Health Organization. Pathologic stages are defined according to the TNM classification.

DNA ploidy, cell cycle analysis, HER-2/neu protein content, p53 protein content, and cathepsin D levels were assessed by immunohistochemical analysis of paraffin embedded tissue. This analysis was performed by an outside reference laboratory with extensive experience in the area of cancer immunohistochemistry (Dianon Laboratories, Stratford, CT) using standard techniques.

Neovascularization in the tumor was evaluated in paraffin-embedded tissue primarily fixed in zinc-buffered formalin. Endothelial cells were stained using antisera against Factor VIII (Dako Polyclonal, Santa Barbara, CA) and the avidin-biotin peroxidase method. Representative areas of the tumor were selected, and microvessel density was assessed using MacMeasure morphometry software (Wayne Rasband, Research Services Branch, NIMH) and a microdigipad (GTCO, Bethesda, MD) with tracings of eighteen x400 photomicrographs of representative areas of each tumor.

Tumor drug metabolism parameters were assessed using fresh tissue. Fresh tumor specimens (at least 500 mg) were minced into small, 2-3 mm pieces and washed twice with cold isotonic saline. Glutathione (GSH) levels were determined in tissue extracts by a specific, and sensitive fluorometric assay using o-phthaladehyde as the fluorescent agent, as described by Hissin and Hilf.³⁷ GSH concentration will be expressed based on milligram protein and milligram DNA.

Patient follow-up. Breast cancer treatment recommendations and treatment

received were recorded for each patient by review of the medical record. Type of surgery was recorded as biopsy, defined as incomplete removal of the tumor for diagnostic purposes only, breast conserving surgery with axillary dissection, simple mastectomy (no axillary dissection), or modified radical mastectomy. Because part of the data was collected retrospectively and the nature of the discussion between the surgeon and patient was not always recorded in the medical record, information regarding the advised surgical approach, in contrast to the ultimate surgical approach, was not available. For the purposes of this analysis, women were considered candidates for breast conservation if their primary tumor was ≤ 4 cm. in size, not fixed to the chest wall or overlying skin, and there was no evidence of distant metastatic disease.

Adjuvant therapy characteristics, including radiation therapy, chemotherapy, and hormonal therapy were recorded for each patient. Radiation therapy factors considered in this analysis included the site of treatment, the total dose received in relation to the total dose planned, and the timing of the treatment.

Chemotherapy received was classified into two general categories: methotrexatebased (in all instances cyclophosphamide, methotrexate, and 5-fluorouracil--CMF), or doxorubicin-based (cyclophosphamide, doxorubicin, +/- 5 fluorouracil--CAF or AC, and in one instance, doxorubicin alone). A modified approach to calculating chemotherapy drug dose intensity was used as a summary descriptor of dose prescribed and received as well as compliance with therapy and completion of recommended chemotherapy course. To calculate dose intensity as used in this report, the method described by Longo, et al. 38 was used with one modification. For subjects who did not complete the prescribed course of therapy, the total time of therapy used in the denominator of the equation was not the actual weeks over which therapy was received, but rather the total number of weeks necessary to complete the entire planned course of chemotherapy. For example, if a subject received only one cycle of a planned six cycles of CMF chemotherapy, the denominator of the dose intensity equation would be 24 weeks, not 4 weeks. If a subject received two cycles of a planned six cycles of chemotherapy, and had a one week treatment delay, the denominator would be 25 weeks, not 9 weeks. This modification was used to allow for the development of a single measure which summarized dose prescribed and received, treatment delays, and compliance with the planned regimen. In all other respects dose intensity was calculated as outlined in the above referenced article.

Because the intent of the analysis was not to compare the different chemotherapy regimens used, but to evaluate the overall quality of the treatment received, the calculated dose intensities were standardized between regimens by dividing the calculated dose intensity by the projected dose intensity,³⁸ yielding a single value for the treatment course, the percent planned dose intensity received by the subject.

Total treatment received for breast cancer was reviewed by one of the investigators, blinded to the race and outcome of the subject, and classified as optimal or suboptimal based upon previously defined criteria. Optimal therapy required the following components: 1) complete surgical resection of the tumor (gross and microscopic) and axillary lymph node dissection was required for all women without distant metastatic disease, 2) radiation therapy was mandatory for all women who underwent breast

conserving surgery and for all women with primary tumors larger than 5 cm. in maximal diameter, 3) adjuvant systemic therapy was required as outlined in the reports from the Fourth and Fifth International Conferences on the Adjuvant Therapy of Primary Breast Cancer, 39,40 except that women with lymph-node negative cancer diagnosed prior to January 1990 were not required to receive adjuvant chemotherapy or hormonal therapy to have their treatment judged as optimal, 4) received chemotherapy dose intensity, calculated as described above, had to be $\geq 85\%$ of the planned dose intensity, and 5) tamoxifen, if prescribed, had to be continued for at least 5 years. Women not meeting the above five criteria were judged to have received suboptimal therapy.

Patient outcome data including response to therapy, time to relapse, and survival were also obtained by review of the medical record.

Statistical analysis. Differences between blacks and whites in categorical breast cancer biologic characteristics will be analyzed by a chi-square test for independence. Continuous variables will be evaluated by the Student's t test and/or Mann-Whitney U test for ranked data. Relapse-free survival and overall survival will be estimated using the Kaplan-Meier product limit method, with differences in survival between black and white patients assessed by a log rank test.

Results

As of May 1, 1997, 140 women have been enrolled in the prospective study. One hundred and fifty additional women have been included in the analysis of clincial presentation and treatment characteristics.

There were 133 black women and 157 white women in the analysis of clinical characteristics at presentation. The demographic characteristics of the total population are described in Table 1. Black women were slightly older, more likely to be covered by Medicare, and less likely to have Medicaid or no health care coverage.

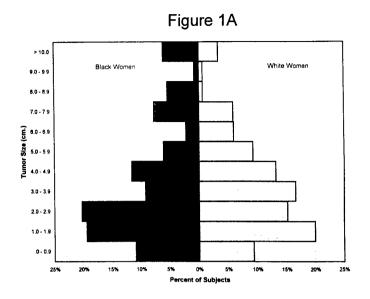
Table 1
Demographic Characteristics

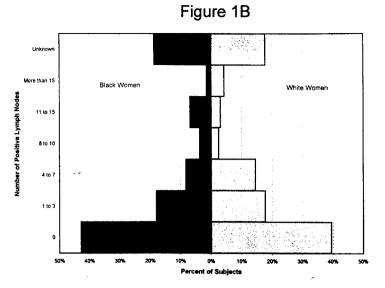
	White (n = 157)	Black (n = 133)	Significance (p value)
Mean age (years)	55.2	58.3	0.04
Post-menopausal (%)	67.5	74.4	0.20
Insurance type (%)			0.03
Medicare	22.5	38.3	
Medicaid	21.2	13.5	
Commercial Insurance	6.6	5.3	
Indigent	49.7	42.9	•

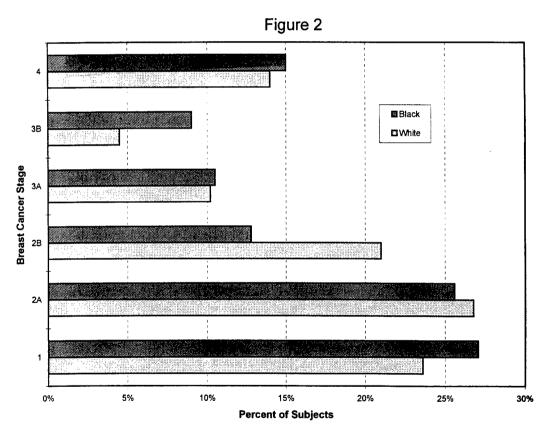
Tumor stage at the time of diagnosis was remarkably similar in black and white women. The mean primary tumor size of black women was 3.77 cm. compared with a mean tumor size of 3.55 for white women. The distribution of tumor size by race (Figure 1A) did not suggest any substantive differences between the two groups, with similar proportions of women presenting with the smallest tumors and similar proportions presenting with very large primary tumors. Likewise, the frequency of cancer spread to the axillary lymph nodes and the number of lymph nodes involved with cancer was very similar between the two groups (Figure 1B). Overall American Joint Committee on Cancer (AJCC) breast cancer stage at diagnosis was not different in the black and white women in this population (Figure 2 and Table 2).

Table 2
Tumor Stage at Diagnosis

	White (n = 157)	Black (n = 133)	Significance (p value)
Tumor Stage (%)			0.84
To	0.6%	0.8%	
T_{1}	32.1%	33.1%	
T_2	40.4%	34.6%	
T ₃	15.4%	16.5%	
T ₄	11.5%	15.0%	
Nodal Stage (%)			0.19
N _o	39.5%	42.1%	
N_1	41.4%	34.6%	
N_2	4.5%	9.0%	
Unknown	14.6%	14.3%	
Metastasis Stage (%)			0.64
M _o	85.0%	86.0%	
M ₁	15.0%	14.0%	
AJCC Stage (%)			0.35
1	23.6%	27.1%	
2A	26.8%	25.6%	
2B	21.0%	12.8%	•
3A	10.2%	10.5%	
3B	4.5%	9.0%	
4	14.0%	15.0%	
Estrogen Receptor (+)	56.3%	48.7%	0.24
Progesterone Receptor (+)	58.7%	44.2%	0.03







Tumor biologic characteristics have been analyzed in the 140 women prospectively enrolled to date. Table 3 below summarizes these results. None of the differences noted between black and white women reach the level of statistical significance (p = 0.05) in univariate analysis at this time.

Table 3
Breast Cancer Biologic Characteristics

	White (n = 69)	Black (n = 71)	Significance (p value)
Estrogen receptor positive	58.6%	49.2%	0.30
Progesterone receptor positive	58.6%	45.9%	0.17
DNA IndexAneuploid	51.5%	41.2%	0.46
S phase fraction high	53.1%	55.3%	0.82
HER-2/neu expression	36.4%	42.6%	0.46
p53 expression	53.0%	48.5%	0.60
Cathepsin D expression	75.8%	76.5%	0.92
Mean microvessel density	13.4	11.5	0.18

The breast cancer primary treatment characteristics of the 290 studied subjects are outlined in Table 4. Overall, 18.3% of the women had breast conservation surgery, with no differences between black and white women. Of the women with tumor sizes ≤ 4 cm., 30.4% of black women and 26.4% of white women underwent breast conservation surgery. Subject age, race, insurance status, and primary tumor size were examined for their association with use of breast conservation. Women choosing lumpectomy with axillary dissection were younger, had smaller primary tumors, and were more likely to have Medicare insurance. Race had no impact on type of surgery received.

Table 3
Breast Cancer Treatment Characteristics

White (n = 157)	Black (n = 133)	Significance (p value)
		0.76
68.2%	64.7%	
18.5%	18.0%	
2.5%	4.5%	
10.8%	12.8%	
39.5%	36.1%	0.53
42.7%	33.1%	0.15
		0.64
52.2%	56.8%	
47.8%	43.2%	
48.4%	52.6%	0.76
	(n = 157) 68.2% 18.5% 2.5% 10.8% 39.5% 42.7% 52.2% 47.8%	(n = 157) (n = 133) 68.2% 64.7% 18.5% 18.0% 2.5% 4.5% 10.8% 12.8% 39.5% 36.1% 42.7% 33.1% 52.2% 56.8% 47.8% 43.2%

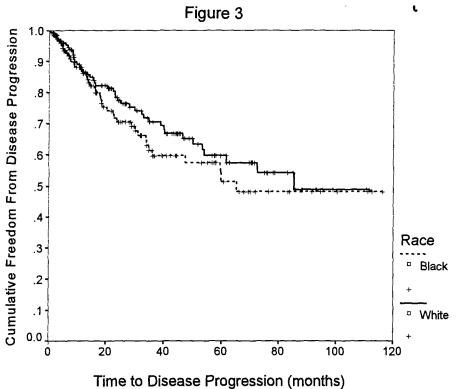
Radiation therapy was used in similar proportions of black and white women.

Overall, 33.8% of women received radiation to their breast or chest wall in the postoperative period, and 4.1% of women received radiation therapy to sites of distant metastases at the time of diagnosis. All women who underwent breast conserving surgery received radiation to their breast.

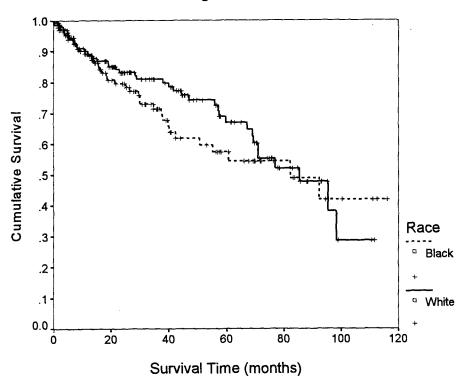
Chemotherapy (either adjuvant therapy or treatment for distant metastases) was given to 42.7% of white women and 33.1% of black women. Approximately half of the women receiving chemotherapy were treated with CMF and half were treated with doxorubicin containing regimens. There was also no difference between black and white women in the percent of planned chemotherapy dose intensity. Black women received a median of 91% of the planned dose intensity with a range of 11-102%; white women received a median of 87% of the planned dose intensity with a range of 8-116%. Forty percent of women received less than 85% of the planned chemotherapy dose intensity. The cause of the low dose intensity was relatively evenly split between dose reductions due to toxicity, patient choice to discontinue chemotherapy, and under-dosing of chemotherapy in obese patients.

Total therapy for primary breast cancer was judged to be optimal in 63.1% of women. Once again, no difference between black and white women was observed. Reasons for suboptimal therapy included receipt of chemotherapy dose intensity less than 85% of that recommended (36%), patient choice not to receive recommended therapy (39%), and failure of the physician to recommend the optimal therapy (25%). Subject age, race, insurance type, tumor size, nodal involvement, presence of distant metastasis, and estrogen receptor levels were evaluated as possible factors associated with receipt of suboptimal therapy. Women who received optimal therapy were more likely to have primary tumors smaller than 5.0 cm (odds ratio = 2.7, 95% confidence interval = 1.5 to 4.8) or to have distant metastatic disease (OR = 2.4, 95% CI = 1.0 to 5.6). Age, race, insurance type, nodal status, and estrogen receptor level did not predict likelihood of optimal treatment.

The median time to disease progression was 5.4 years in black women and 7.1 years in white women. While the difference seems substantial, inspection of the graphical representation of the data demonstrates very similar progression risks for black and white women (Figure 3, log-rank p = 0.35). Overall survival was also very similar between black and white women (Figure 4). The median survival of the black women in the study was 6.9 years and the median survival of the white women studied was 7.1 years. Cox regression analysis identified a hazard rate for death among black women to be 1.24 (95% confidence interval 0.80 to 1.92) compared to white women. Adjustment for commonly accepted prognostic factors, including tumor size, lymph node metastasis, distant metastatic disease, estrogen receptor status, receipt of optimal therapy, and age, did not substantively change the hazard ratio or narrow the confidence interval.







In both racial groups, breast cancer was the primary cause of death in approximately 75% of the women. There were two possible treatment related deaths, a women suffering a post-operative myocardial infarction following modified radical mastectomy and a women developing a myelodysplastic syndrome five years after adjuvant CMF. The remainder of the women studied died from unrelated causes. Specifically, there did not appear to be a higher mortality from comorbid conditions in black women.

Conclusions

To date, patient accrual and sample processing has gone very smoothly. The number of women enrolled in the study over the last nine months has been slightly less than expected. This is the result of an increasingly common practice at this institution to treat some women with locally advanced breast cancer with neoadjuvant chemotherapy. Most often, the diagnosis of breast cancer in these women has been made by aspiration biopsy or core needle biopsy. The amount of tissue obtained with these techniques is generally insufficient to perform the planned biochemical analyses. While subsequent surgical excision of the tumor often yields residual cancer, we had made the decision when developing this study not to analyze tumor that had been exposed to previous chemotherapy, as the effects of this exposure on the characteristics of the tumor were unknown. It is, therefore, likely, given the present subject accrual rates, that continued patient enrollment will need to extend into the latter half of 1998 to meet the enrollment goal of 200 women.

Our study, focusing only on women of lower socioeconomic status did not reveal any tumor stage differences between black and white women. In addition, the pattern of stage distribution in this racially mixed population did demonstrate a shift towards more advanced disease, very similar to that typically described in black women. Only 40% of the women in this study were node-negative, compared to the SEER data in which 50% of women were node-negative. Likewise, we observed a higher incidence of distant metastatic disease at the time of diagnosis than is typically recognized (14% in the current study compared to 7% in the SEER data).

To date, we have identified no significant racial differences in the biological characteristics of breast cancer. It is, however, too early to conclude that a difference does not exist. The number of women studied is too small to say confidently that the tumors in black and white women are biologically similar. Firm conclusions in this area will need to await completion of the study.

Consistent with the more recent reports, we found no difference between black and white women in the type of surgery performed for the treatment of the primary breast lesion. Eighteen percent of all women and 28% of women with tumors ≤ 4 cm. had breast conservation. A relatively high number of women did not have definitive surgical treatment of the breast cancer. Eleven percent of women had a biopsy only. The large majority of these women had documented distant metastatic disease, although there were several

women with earlier stage tumors who refused further treatment after the diagnosis of the cancer.

We, likewise, found no substantial racial difference in the frequency with which radiation therapy to the breast or chest wall was employed in the treatment regimen (36% in black women and 39% in white women). Of note, all women in this study who had breast conserving surgery underwent a complete course of post-operative breast irradiation.

We found that white women were slightly more likely to receive chemotherapy than black women, although this difference was not statistically significant. Similar proportions of black and white women received doxorubicin-based chemotherapy. In addition, the proportion of optimal dose intensity received by black and white women was also very similar (91% of planned dose intensity in black women and 87% of planned dose intensity in white women). Reasons for receiving lower than planned chemotherapy dose intensity were not different in black and white women, with approximately one-third experiencing treatment delays or dose reductions as a result of chemotherapy toxicity, one-third receiving a suboptimal chemotherapy dose for their actual body weight (this was most commonly seen in obese women), and one-third refusing to complete the entire planned course of chemotherapy.

While examining the proportion of women that receive the different treatment modalities provides some insight into possible variations in the care of women with breast cancer, this approach does not directly assess the primary issue of treatment quality. Treatment standards for breast cancer are dependent upon age of the patient, stage of disease, and hormone receptor status. In order to accurately determine if racial differences in breast cancer treatment exist, these factors must be considered. We attempted to incorporate this issue into the current study by calculating the proportion of women that received "optimal" therapy for breast cancer. The recommendations developed by the Fourth and Fifth International Conferences on the Adjuvant Therapy of Breast Cancer^{39,40} were used as a guide for our definition of patient specific "optimal" therapy. In addition to the published suggestions, we included requirements for breast or chest wall irradiation for women with breast conservation surgery or large primary tumors. Minimum acceptable chemotherapy dose intensity guidelines for all women receiving adjuvant chemotherapy were also incorporated into the definition of optimal therapy. 41,42

Using these guidelines, we found that 63% of white women and 64% of black women received optimal therapy for their breast cancer. While it is clear that no racial difference in quality of care, as defined by our guidelines, exists, it was disappointing to see that fully one third of women in this cohort received care judged to be suboptimal. Demographic factors did not appear to influence the likelihood of receiving suboptimal therapy.

It is impossible to determine if socioeconomic factors were at all important in determining quality of care. There have been no previously reported rigorous studies of breast cancer treatment quality in an undifferentiated cohort of women more representative of the general population of the United States. A recent study evaluating management of local/regional breast cancer in women insured by Blue Cross/Blue Shield found that only

54% of women under age 51 received adjuvant chemotherapy, suggesting that problems with delivery of optimal therapy exist not only among women served at the public hospitals but also among women with commercial health care insurance.⁴³ Further studies in this area would certainly be helpful in understanding the relationship between socioeconomic status and quality of cancer care.

The only characteristics identified which did change the risk of receiving suboptimal therapy were characteristics related to disease stage. Women with tumors larger than five centimeters were less likely to received optimal therapy, primarily because they were not routinely referred for chest wall irradiation. In a similar manner, women with metastatic disease were more likely to receive optimal therapy, primarily because not all women with local or regional disease were referred to medical oncology for adjuvant systemic therapy. Physician-related issues regarding quality of care only accounted for 33% of the women who received suboptimal treatment, however. The remaining cases of suboptimal care were a result of patient choice not to receive recommended treatment or treatment toxicity prohibiting optimal therapy delivery.

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DEPARTMENT OF THE ARMY

US ARMY MEDICAL RESEARCH AND MATERIEL COMMAND 504 SCOTT STREET FORT DETRICK, MARYLAND 21702-5012

REPLY TO ATTENTION OF:

MCMR-RMI-S (70-1y)

3 JUN 2001

MEMORANDUM FOR Administrator, Defense Technical Information Center (DTIC-OCA), 8725 John J. Kingman Road, Fort Belvoir, VA 22060-6218

SUBJECT: Request Change in Distribution Statement

- 1. The U.S. Army Medical Research and Materiel Command has reexamined the need for the limitation assigned to technical reports. Request the limited distribution statement for reports on the enclosed list be changed to "Approved for public release; distribution unlimited." These reports should be released to the National Technical Information Service.
- 2. Point of contact for this request is Ms. Judy Pawlus at DSN 343-7322 or by e-mail at judy.pawlus@dat.amedd.army.mil.

FOR THE COMMANDER:

Encl

PHYLIS M. RINEHART

Deputy Chief of Staff for Information Management

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